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FORM PTO-1449/A and B (Modified)

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Sheet

INFORMATION DISCLOSURES
STATEMENT BY APPLICANT

APPLICATION NO.: 10/821,811

ATTY. DOCKET NO.: P0453.70115US01

FILING DATE:

April 8, 2004

CONFIRMATION NO.: Not Yet Assigned

APPLICANT:

Sanghvi et al.

of 6

GROUP ART UNIT: Not Yet Assigned

EXAMINER: Not Yet Assigned

U.S. PATENT DOCUMENTS

Examiner's	Cite	U.S. Patent Doo	cument	Name of Patentee or Applicant of Cited	Date of Publication or of issue	
Initials	No.	Number	Kind Code	Document	of Cited Document MM-DD-YYYY	
	A1	2001/0018413	A1	Crain, et al.	08-30-2001	
	A2	2002/0028825	A1	Foss, et al.	03-07-2002	
	A3	2001/0033865	A1	Oshlack, et al.	10-25-2001	
	A4	2001/0036476	A1	Oshlack, et al.	11-01-2001	
	A5	2001/0047005	A1	Farrar, et al.	11-29-2001	
	A6	4,176,186		Goldberg, et al.	11-27-1979	
	A7	4,719,215		Goldberg	01-12-1988	
	A8	4,861,781		Goldberg	08-29-1989	
	A9	4,987,136		Kreek, et al.	01-22-1991	
	A10	5,102,887		Goldberg	04-07-1992	
	A11	5,270,328		Cantrell, et al.	12-14-1993	
	A12	5,472,943		Crain, et al.	12-05-1995	
	A13	5,512,578		Crain, et al.	04-30-1996	
	A14	5,767,125		Crain, et al.	06-16-1998	
	A15	5,811,451		Minoia, et al.	09-22-1998	
	A16	5,866,164		Kuczynski, et al.	02-02-1999	
	A17	5,958,452		Oshlack, et al.	09-28-1999	
	A18	5,972,954		Foss, et al.	10-26-1999	
	A19	6,096,756		Crain, et al.	08-01-2000	
	A20	6,194,382	B1	Crain, et al.	02-27-2001	
	A21	6,261,599	B1	Oshlack, et al.	07-17-2001	
	A22	6,274,591	B1	Foss, et al.	08-14-2001	
	A23	6,395,705	B2	Crain, et al.	05-28-2002	
	A24	6,419,959	B1	Walter, et al.	07-16-2002	
	A25	6,451,806	B2	Farrar	09-17-2002	
	A26	6,559,158	B1	Foss, et al.	05-06-2003	
	A27	6,608,075	B1	Foss, et al.	08-19-2003	
	A28	RE36,547		Crain, et al.	02-01-2000	
	A29	2002/0188005	A1	Farrar, et al.	12-12-2002	

FOREIGN PATENT DOCUMENTS

Examiner's	Cite	Foreign Patent Document		Foreign Patent Document		Consider Dotage Document I Name of Patentee of Applicant of Cited I	Foreign Patent Document Name of Pat		Date of Publication of	Translation
Initials	No.	Office/ Country	Number	Kind Code	(not necessary)	Cited Document MM-DD-YYYY	(Y/N)			
	B1	AU	610,561	_	Shelley	08-17-1988				
	B2	CA	1,315,689		The University of Chicago	04-06-1993				
	B3	EP	0278821	A1	Shelly (Abstract)	08-17-1988				

ALL REFERENCES CONSIDERED EXCEPT WHERE LINED THROUGH. /D.J./ (09/21/2008)

FORM PTO-1449/A and B (Modified) INFORMATION DISCLOSURE STATEMENT BY APPLICANT Sheet 2 of 6 APPLICATION NO.: 10/821,811 ATTY. DOCKET NO.: P0453.70115US01 APPLICATION NO.: 10/821,811 ATTY. DOCKET NO.: P0453.70115US01 FILING DATE: April 8, 2004 CONFIRMATION NO.: Not Yet Assigned APPLICANT: Sanghvi et al. GROUP ART UNIT: Not Yet Assigned EXAMINER: Not Yet Assigned

	B4	EP	0352361	A 1	The Rockefeller University	01-31-1990	
	B5	EP	278,821	A 1	Marc Yves Shelly (Derwent Abstract)	08-17-1988	•
	B6	EP	306,575	B 1	The Univ. of Chicago	03-15-1989	
•	B7	EP	352,361	A 1	The Rockefeller University	01-31-1990	
	B8	EP	760,661	B 1	Minoia, et al.	12-30-1998	
_	B9	JP	2,625,457	B2	Goldberg (Derwent Abstract)	07-02-1997	
	B10	NZ	222,911		The Univ. of Chicago	12-14-1987	
	B11	wo	83/03197	A 1	The Rockefeller University	09-29-1983	
	B12	wo	88/05297	A 1	Shelly	07-28-1988	
	B13	wo	95/31985	A2	Minoia, et al.	11-30-1995	
	B14	wo	97/33566		Alza Corp.	09-18-1997	
	B15	wo	98/25613		Klinge Pharma GmbH	06-18-1998	Yes
	B16	wo	01/13909	A2	Critical Care Pharm.	03-01-2001	
	B17	wo	01/37785	A2	Adolor Corp.	05-31-2001	
	B18	wo	01/41705	A2	Adolor Corp.	06-14-2001	
	B19	wo	01/42207	A2	Adolor Corp.	06-14-2001	
	B20	wo	01/85257	A2	Pain Therapeutics, Inc	11-15-2001	
	B21_	wo	02/060870	A2	Adolor Corp.	08-08-2002	

OTHER ART — NON PATENT LITERATURE DOCUMENTS

Examiner's	Cite	Include name of the author (in CAPITAL LETTERS) title of the article (when appropriate), title of the item	Translation	
Initials	No	(book, magazine, journal, serial, symposium, catalog, etc.), date, relevant page(s), volume-issue number(s),	(Y/	N)
		publisher, city and/or country where published.	<u> </u>	
	04	AKINBAMI et al., Effect of a peripheral and a central acting opioid antagonist on the testicular response to]	
	C1	stress in rats. Neuroendocrinology. 1994 Apr;59(4):343-8.	 	
	00	AMIN et al., Efficacy of methylnaltrexone versus naloxone for reversal of morphine-induced depression of		
	C2	hypoxic ventilatory response. Anesth Analg. 1994 Apr;78(4):701-5.	ļ	
		AMIR, Naloxone improves, and morphine exacerbates, experimental shock induced by release of endogenous		
	C3	histamine by compound 48/80. Brain Res. 1984 Apr 9;297(1):187-90.		
	C4	AMIR et al., Endorphins in endotoxin-induced hyperglycemia in mice. Arch Toxicol Suppl. 1983;6:261-5.		
		ARGENTIERI et al., Interaction of the opiate antagonist, naltrexone methyl bromide, with the acetylcholine		
	C5	receptor system of the motor end-plate. Brain Res. 1983 Oct 31;277(2):377-9.		
		BARATTI et al., Brain opioid peptides may participate in the reversal of pentylenetetrazol-induced amnesia.		
	C6	Methods Find Exp Clin Pharmacol. 1990 Sep;12(7):451-6.		
		BEDINGFIELD et al., Methylnaltrexone attenuates taste aversion conditioned by low-dose ethanol. Alcohol.		
	C7	1998 Jan;15(1):51-4.		
		BIANCHETTI et al., Quaternary derivatives of narcotic antagonists: stereochemical requirements at the chiral		-
	C8	nitrogen for in vitro and in vivo activity. Life Sci. 1983;33 Suppl 1:415-8.		
		BIANCHI et al., Quaternary narcotic antagonists' relative ability to prevent antinociception and gastrointestinal		
		transit inhibition in morphine-treated rats as an index of peripheral selectivity. Life Sci. 1982 May		
	C9	31;30(22):1875-83.		
		BICKEL, Stimulation of colonic motility in dogs and rats by an enkephalin analogue pentapeptide. Life Sci.		
	C10	1983;33 Suppl 1:469-72.		
		BLANK et al., Central, stereoselective receptors mediate the acute effects of opiate antagonists on luteinizing		
	C11	hormone secretion. Life Sci. 1986 Oct 27;39(17):1493-99.		
···		BRIX-CHRISTENSEN et al., Endogenous morphine is produced in response to cardiopulmonary bypass in	† †	
	C12	neonatal pigs. Acta Anaesthesiol Scand. 2000 Nov;44(10):1204-8.	1	
· · · · · · ·		BROWN et al., Opiate antagonists: central sites of action in suppressing water intake of the rat. Brain Res. 1981		
	C13	Sep 28;221(2):432-6.		

FORM PTO-1449/A and B (Modified)	APPLICATION NO.	: 10/821,811	ATTY. DOCKET NO.: P0453.70115US01
INFORMATION DISCLOSURE	FILING DATE:	April 8, 2004	CONFIRMATION NO.: Not Yet Assigned
STATEMENT BY APPLICANT	APPLICANT:	Sanghvi et al.	
	GROUP ART UNIT:	Not Vet Assigned	EXAMINER: Not Yet Assigned
Sheet 3 of 6	Choo! ART ONT.	110t 10t Assigned	DATIVITYDIC. 1100 1 of Assigned

		BROWN et al., Reversal of morphine-induced catalepsy in the rat by narcotic antagonists and their quaternary		
	C14	derivatives. Neuropharmacology. 1983 Mar;22(3):317-21.		
		BROWN et al., The use of quaternary narcotic antagonists in opiate research. Neuropharmacology. 1985		
	C15	Mar;24(3):181-91. Review.		
		CALCAGNETTI et al., Quaternary naltrexone reveals the central mediation of conditional opioid analgesia.		
	C16	Pharmacol Biochem Behav. 1987 Jul;27(3):529-31.		
		CHANG et al., An antiabsorptive basis for precipitated withdrawal diarrhea in morphine-dependent rats. J		
	C17	Pharmacol Exp Ther. 1984 Feb;228(2):364-9.		
		CULPEPPER-MORGAN et al., Treatment of opioid-induced constipation with oral naloxone: a pilot study. Clin		
	C18	Pharmacol Ther. 1992 Jul;52(1):90-5 (ABSTRACT ONLY).		
	040	EISENBERG, Effects of naltrexone on plasma corticosterone in opiate-naive rats: a central action. Life Sci.		
	C19	1984 Mar 19;34(12):1185-91.		
	Can	FERNANDEZ-TOME et al., Interaction between opioid agonists or naloxone and 5-HTP on feeding behavior in		
	C20	food-deprived rats. Pharmacol Biochem Behav. 1988 Feb;29(2):387-92.		
	C21	FOSS, A review of the potential role of methylnaltrexone in opioid bowel dysfunction. Am J Surg. 2001	.	
	021	Nov;182(5A Suppl):19S-26S. Review. FOSS et al., 1995 Annual scientific meeting of the American Society of Anesthesiologists. Atlanta, Georgia,		
	C22	October 21-25, 1995. Abstracts. Anesthesiology. 1995 Sep;83(3A Suppl):A361.		
	022	FOSS et al., Prevention of apomorphine- or cisplatin-induced emesis in the dog by a combination of		
	C23	methylnaltrexone and morphine. Cancer Chemother Pharmacol. 1998;42(4):287-91.	.	
	023	FOSS et al., Safety and tolerance of methylnaltrexone in healthy humans: a randomized, placebo-controlled,		
1	C24	intravenous, ascending-dose, pharmacokinetic study. J Clin Pharmacol. 1997 Jan;37(1):25-30.		
	<u> </u>	FOSS et al., Dose-related antagonism of the emetic effect of morphine by methylnaltrexone in dogs. J Clin		
	C25	Pharmacol. 1993 Aug;33(8):747-51.		
		FOSS et al., Effects of methylnaltrexone on morphine-induced cough suppression in guinea pigs. Life Sci.		
	C26	1996;59(15):PL235-8.		
		FOSS et al., Methylnaltrexone reduces morphine-induced postoperative emesis by 30%. Anesth Analg.		
	C27	1994;78:S119.		
		FRANCE et al., Comparison of naltrexone and quaternary naltrexone after systemic and intracerebroventricular		
	C28	administration in pigeons. Neuropharmacology. 1987 Jun;26(6):541-8.		
		FRANCE et al., Intracerebroventricular drug administration in pigeons. Pharmacol Biochem Behav. 1985		
	C29	Nov;23(5):731-6.		
		FRIEDMAN et al., Opioid antagonists in the treatment of opioid-induced constipation and pruritus. Ann		
	C30	Pharmacother. 2001 Jan;35(1):85-91. Review.		
	004	GMEREK et al., Independent central and peripheral mediation of morphine-induced inhibition of	. 1	
	C31	gastrointestinal transit in rats. J Pharmacol Exp Ther. 1986 Jan;236(1):8-13.		
	C32	HEIN et al., Pharmacological analysis of the discriminative stimulus characteristics of ethylketazocine in the	. 1	
		rhesus monkey. J Pharmacol Exp Ther. 1981 Jul;218(1):7-15.		
1	C33	HOWD et al., Naloxone and intestinal motility. Experientia. 1978 Oct 15;34(10):1310-1.		
	C34	JALOWIEC et al., Suppression of juvenile social behavior requires antagonism of central opioid systems.	ı İ	
	034	Pharmacol Biochem Behav. 1989 Jul;33(3):697-700. JANKOVIC et al., Quaternary naltrexone: its immunomodulatory activity and interaction with brain delta and		
	C35	kappa opioid receptors. Immunopharmacology. 1994 Sep-Oct;28(2):105-12.		
	000	KAUFMAN et al., Role of opiate receptors in the regulation of colonic transit. Gastroenterology. 1988		
	C36	Jun;94(6):1351-6.		
		KIM et al., Assay for methylnaltrexone in rat brain regions and serum by high-performance liquid		
	C37	chromatography with coulometric electrochemical detection. Chromatographia. 1989 Oct;28(7-8):359-63.		
		KINSMAN et al., Effect of naloxone on feedback regulation of small bowel transit by fat. Gastroenterology.		
	C38	1984 Aug;87(2):335-7.		
		KOBLISH et al., Behavioral profile of ADL 8-2698, a novel GI-restricted μ opioid receptor antagonist. Society		
	C39	for Neuroscience Abstracts. 2001;27(2):2407.		
		KOBYLECKI et al., N-Methylnalorphine: definition of N-allyl conformation for antagonism at the opiate		
	C40	receptor. J Med Chem. 1982 Nov;25(11):1278-80.	igsquare	
	C41	KOCZKA, et al., Acta Chimica Academica Scien. Hung. (1967) 51(4), 393-02		

FORM PTO-1449/A and B (Modified)	APPLICATION NO.: 10	0/821,811	ATTY. DOCKET NO.: P0453.70115US01
INFORMATION DISCLOSURE	FILING DATE: A	pril 8, 2004	CONFIRMATION NO.: Not Yet Assigned
STATEMENT BY APPLICANT	APPLICANT: Sa	anghvi et al.	
Sheet 4 of 6	GROUP ART UNIT: N	ot Yet Assigned	EXAMINER: Not Yet Assigned

		KOOB et al., Effects of opiate antagonists and their quaternary derivatives on heroin self-administration in the	
	C42	rat. J Pharmacol Exp Ther. 1984 May;229(2):481-6.	
	1072	KOTAKE et al., Variations in demethylation of N-methylnaltrexone in mice, rats, dogs, and humans.	
	C43	Xenobiotica. 1989 Nov;19(11):1247-54.	
•	- 040	KROMER et al., Endogenous opioids, the enteric nervous system and gut motility. Dig Dis. 1990;8(6):361-73.	-
	C44	Review.	
	 	KROMER et al., The current status of opioid research on gastrointestinal motility. Life Sci. 1989;44(9):579-89.	
	C45	Review.	1
	C46	LEANDER, A kappa opioid effect: increased urination in the rat. J Pharmacol Exp Ther. 1983 Jan;224(1):89-94.	+
	C47	LITTLE, et al., Society for Neuroscience Abstracts, 27 (2); 2001, p. 2407	
	C48	LIVINGSTON et al., Postoperative ileus. Dig Dis Sci. 1990 Jan;35(1):121-32. Review.	
	C49	LYDON et al., ESA Free Paper Prize Competition. Eur J Anaesthesiol. 2001 Apr;18 Suppl 21:92.	
		LYSLE et al., Modulation of immune status by a conditioned aversive stimulus: evidence for the involvement of	
	C50	endogenous opioids. Brain Behav Immun. 1992 Jun;6(2):179-88.	
		MAGNAN et al., The binding spectrum of narcotic analgesic drugs with different agonist and antagonist	
	C51	properties. Naunyn Schmiedebergs Arch Pharmacol. 1982 Jun;319(3):197-205.	- 1
	C52	MANARA, et al., Adv. Endog. Exog. Opioids, Poroc. Int. Narc. Res. Conf., 12th (1981), 402-4	
		MANARA et al., The central and peripheral influences of opioids on gastrointestinal propulsion. Annu Rev	
	C53	Pharmacol Toxicol. 1985;25:249-73. Review.	
		MICKLEY et al., Quaternary naltrexone reverses morphine-induced behaviors. Physiol Behav. 1985	
	C54	Aug;35(2):249-53.	
		MISRA et al., Intravenous kinetics and metabolism of [15,16-3H]naltrexonium methiodide in the rat. J Pharm	
	C55	Pharmacol. 1987 Mar;39(3):225-7.	
		MOERMAN et al., Evaluation of methylnaltrexone for the reduction of postoperative vomiting and nausea	
	C56	incidences. Acta Anaesthesiol Belg. 1995;46(3-4):127-32.	
	C57	MOSS, et al., N. Engl. J. Med., (2002) 346 (6), 455	
		MUCHA, Is the motivational effect of opiate withdrawal reflected by common somatic indices of precipitated	
	C58	withdrawal? A place conditioning study in the rat. Brain Res. 1987 Aug 25;418(2):214-20.	
		MUCHA, Taste aversion involving central opioid antagonism is potentiated in morphine-dependent rats. Life	_
	C59	Sci. 1989;45(8):671-8.	
	C60	MURPHY et al., Anesthesiology, Sept. (1999), 91 (3A) p. A349 (Abstract)	
		MURPHY et al., Pharmacokinetic profile of epidurally administered methylnaltrexone, a novel peripheral opioid	
	C61	antagonist in a rabbit model. Br J Anaesth. 2001 Jan;86(1):120-2.	
	 •••	MURPHY et al., American Society of Anesthesiologists 1999 annual meeting. Dallas, Texas, USA. October 9-	
	C62	13, 1999. Abstracts. Anesthesiology. 1999 Sep;91(3A Suppl):A349.	
		MURPHY et al., Opioid-induced delay in gastric emptying: a peripheral mechanism in humans. Anesthesiology.	-
	C63	1997 Oct;87(4):765-70.	1
		MURPHY et al., Opioid antagonist modulation of ischaemia-induced ventricular arrhythmias: a peripheral	
	C64	mechanism. J Cardiovasc Pharmacol. 1999 Jan;33(1):122-5.	1
		NARANJO et al., Evidence for a central but not adrenal, opioid mediation in hypertension induced by brief	
	C65	isolation in the rat. Life Sci. 1986 May 26;38(21):1923-30.	
000000000000000000000000000000000000000		NELSON, Dissertation Abstracts International, (62/03-B), p. 1635 (Abstract)	
	- 000	ODIO et al., Central but not peripheral opiate receptor blockade prolonged pituitary-adrenal responses to stress.	
	C67	Pharmacol Biochem Behav. 1990 Apr;35(4):963-9.	
	 55.	OSINSKI et al., Determination of methylnaltrexone in clinical samples by solid-phase extraction and high-	
		performance liquid chromatography for a pharmacokinetics study. J Chromatogr B Analyt Technol Biomed Life	
	C68	Sci. 2002 Nov 25;780(2):251-9.	
	1	PAPPAGALLO, Incidence, prevalence, and management of opioid bowel dysfunction. Am J Surg. 2001	-+
	C69	Nov;182(5A Suppl):11S-18S. Review.	ļ
	555	POLAK et al., Enkephalin-like immunoreactivity in the human gastrointestinal tract. Lancet. 1977 May	-
	C70	7;1(8019):972-4.	
		POWELL et al., Paradoxical effects of the opioid antagonist naltrexone on morphine analgesia, tolerance, and	
	C71	reward in rats. J Pharmacol Exp Ther. 2002 Feb;300(2):588-96.	

FORM PTO	-1449/A and B (M	lodifie	d)	APPLICATION NO.:	10/821,811	ATTY. DOCKET NO.: P0453.70115US01
	RMATION D			FILING DATE:	April 8, 2004	CONFIRMATION NO.: Not Yet Assigned
STAT	EMENT BY	APP	LICANT	APPLICANT:	Sanghvi et al.	
				GROUP ART UNIT:	Not Yet Assigned	EXAMINER: Not Yet Assigned
Sheet	5	of	6			

	072	QUOCK et al., Narcotic antagonist-induced hypotension in the spontaneously hypertensive rat. Life Sci. 1985	
	C73	Sep 2;37(9):819-26.	
	C74	QUOCK et al., Narcotic antagonist potentiation of apomorphine drug effect: a stereospecific, centrally mediated drug action. Prog Neuropsychopharmacol Biol Psychiatry. 1985;9(3):239-43.	
•	C75	RAMABADRAN, Effects of N-methylnaloxone and N-methylnaltrexone on nociception and precipitated abstinence in mice. Life Sci. 1982 Sep 20-27;31(12-13):1253-6.	
	C76	RIVIÈRE et al., Fedotozine reverses ileus induced by surgery or peritonitis: action at peripheral kappa-opioid receptors. Gastroenterology. 1993 Mar;104(3):724-31.	
	C77	ROBINSON et al., Oral naloxone in opioid-associated constipation. Lancet. 1991 Aug 31;338(8766):581-2.	
	C78	ROGER et al., Colonic motor responses in the pony: relevance of colonic stimulation by opiate antagonists. Am J Vet Res. 1985 Jan;46(1):31-5.	
	C79	RUSSELL et al., Antagonism of gut, but not central effects of morphine with quaternary narcotic antagonists. Eur J Pharmacol. 1982 Mar 12;78(3):255-61.	
	C80	SCHAEFER et al., Effects of opioid antagonists and their quaternary derivatives on locomotor activity and fixed ratio responding for brain self-stimulation in rats. Pharmacol Biochem Behav. 1985 Nov;23(5):797-802.	
	C81	SCHANG et al., Beneficial effects of naloxone in a patient with intestinal pseudoobstruction. Am J Gastroenterol. 1985 Jun;80(6):407-11.	
	C82	SCHANG et al., How does morphine work on colonic motility? An electromyographic study in the human left and sigmoid colon. Life Sci. 1986 Feb 24;38(8):671-6.	
	C83	SCHILLER et al., Studies of the mechanism of the antidiarrheal effect of codeine. J Clin Invest. 1982 Nov;70(5):999-1008.	
	C84	SCHMIDHAMMER, et al., Helv. Chim. Acta (1994), Vol. 77, No. 6, p. 1585-9	
	C85	SCHMIDHAMMER, et al., Helv. Chim. Acta. (1993) No. 1, p. 476-80	
	C86	SCHOLZ, 2000, 63 (6) p. 103	
	C87	SCHREIER et al., Central regulation of intestinal function: morphine withdrawal diarrhea. Proc West Pharmacol Soc. 1982;25:151-4.	
		SOLVASON et al., Naltrexone blocks the expression of the conditioned elevation of natural killer cell activity in	
	C88	BALB/c mice. Brain Behav Immun. 1989 Sep;3(3):247-62.	
	C89	SWAN, et al., AIDS Research, NIDA Notes, (1995), 10(3), 1-6	
	C90	SYKES, Oral naloxone in opioid-associated constipation. Lancet. 1991 Jun 15;337(8755):1475.	
	C91	TAGUCHI et al., Selective postoperative inhibition of gastrointestinal opioid receptors. N Engl J Med. 2001 Sep 27;345(13):935-40.	
	C92	THOMPSON et al., Opioid stimulation in the ventral tegmental area facilitates the onset of maternal behavior in rats. Brain Res. 1996 Dec 16;743(1-2):184-201.	
	C93	UKAI et al., Suppression of deprivation-induced water intake in the rat by opioid antagonists: central sites of action. Psychopharmacology (Berl). 1987;91(3):279-84.	
	C94	VALENTINO et al., Quaternary naltrexone: evidence for the central mediation of discriminative stimulus effects of narcotic agonists and antagonists. J Pharmacol Exp Ther. 1981 Jun;217(3):652-9.	
-	C95	VALENTINO et al., Receptor binding, antagonist, and withdrawal precipitating properties of opiate antagonists. Life Sci. 1983 Jun 20;32(25):2887-96.	
	C96	WALKER, et al., Psychopharmacology (1991), 104(2), p. 164-6	
		WARREN et al., Effects of quaternary naltrexone and chlordiazepoxide in squirrel monkeys with enhanced	
	C97	sensitivity to the behavioral effects of naltrexone. J Pharmacol Exp Ther. 1985 Nov;235(2):412-7.	$-\!\!\!\!+$
	C98	WILLETTE, et al., Res. Commun. Subst. Abuse (1983), 4(4), 325-37	
	C99	YUAN et al., Drug Dev. Res. (2000) 50(2), 133-141	
	C100	YUAN et al., Gastric effects of methylnaltrexone on mu, kappa, and delta opioid agonists induced brainstem unitary responses. Neuropharmacology. 1999 Mar;38(3):425-32.	
	C101	YUAN et al., Anesthesiology, Sept. (1995), 83 (3A), p A358 (Abstract)	
	C102	YUAN et al., Anesthesiology, Sept. (1995), 83 (3A), p A360 (Abstract)	
	C102	YUAN et al., Anesthesiology, Sept. (1999), 91 (3A) p. A973 (Abstract)	
	0103	YUAN et al., Effects of enteric-coated methylnaltrexone in preventing opioid-induced delay in oral-cecal transit	-+
	C104	time. Clin Pharmacol Ther. 2000 Apr;67(4):398-404.	
	0405	YUAN et al., The safety and efficacy of oral methylnaltrexone in preventing morphine-induced delay in oral-	
	C105	cecal transit time. Clin Pharmacol Ther. 1997 Apr;61(4):467-75. BENCES CONSIDERED EXCEPT WHERE LINED THROUGH. /D.L/ (09/21/2008)	

FORM PTO-1449/A and B (Modified)	APPLICATION NO.:	10/821,811	ATTY. DOCKET NO.: P0453.70115US01
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'STATEMENT BY APPLICANT	APPLICANT:	Sanghvi et al.	
Sheet 6 of 6	GROUP ART UNIT:	Not Yet Assigned	EXAMINER: Not Yet Assigned

:	C106	YUAN et al., Methylnaltrexone prevents morphine-induced delay in oral-cecal transit time without affecting analgesia: a double-blind randomized placebo-controlled trial. Clin Pharmacol Ther. 1996 Apr;59(4):469-75.	
	C107	YUAN, et al., Clinical Pharmacology & Therapeutics (1995) 57(2), p. 138	
•	C108	YUAN et al., Efficacy of orally administered methylnaltrexone in decreasing subjective effects after intravenous morphine. Drug Alcohol Depend. 1998 Oct 1;52(2):161-5.	
	C109	YUAN et al., Effects of methylnaltrexone on morphine-induced inhibition of contraction in isolated guinea-pig ileum and human intestine. Eur J Pharmacol. 1995 Mar 24;276(1-2):107-11.	
	C110	YUAN et al., Effects of subcutaneous methylnaltrexone on morphine-induced peripherally mediated side effects: a double-blind randomized placebo-controlled trial. J Pharmacol Exp Ther. 2002 Jan;300(1):118-23.	
	C111	YUAN et al., Oral methylnaltrexone for opioid-induced constipation. JAMA. 2000 Sep 20;284(11):1383-4.	
	C112	YUAN et al., Methylnaltrexone for reversal of constipation due to chronic methadone use: a randomized controlled trial. JAMA. 2000 Jan 19;283(3):367-72.	
	C113	YUAN et al., Effects of intravenous methylnaltrexone on opioid-induced gut motility and transit time changes in subjects receiving chronic methadone therapy: a pilot study. Pain. 1999 Dec;83(3):631-5.	
	C114	YUAN et al., Effects of methylnaltrexone on chronic opioid induced gut motility and transit time changes. Br J Anaesth. 1998;81(1):94.	
	C115	YUAN et al., Effects of methylnaltrexone on chronic opioid-induced gut motility and transit time changes. University of Leicester – Abstracts from the Eighth International Symposium on Pain, Anaesthesia and	
	C115 C116	Endocrinology. 1997 September 18-19th. [No Author Listed] Oncology. 1996;10(12):1880.	
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ALL REFERENCES CONSIDERED EXCEPT WHERE LINED THROUGH. /D.J./ (09/21/2008)

EXAMINER	/Donna Jagoe/ (9/21/2008)	DATE CONSIDERED

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